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Applicant(s): John Bertin

NOVEL MOLECULES OF THE CARD-RELATED PROTEIN
FAMILY AND USES THEREOF

1 GTCGACCCACGCGTCCGGCAGCAGGCAGGCTGCAGCAGGCGAGCAGCAGCAAGAGTAAAAGG
CAGCTGGGTGCGCAGGCCGTCGTCGGTCCGACGTCGTCGGCTCGTCGTCGTTCTCATTTTCC

63 TGACCGCGGCTGCCCCACCCAGAGCCATGGGGCGGGCACGAGATGCCATCCTGGACGCTCTT
ACTGGCGCCGACGGGTGGGGTCTCGGTACCCCGCCCGTGCTCTACGGTAGGACCTGCGAGAA

1 M G R A R D A I L D A L

125 GAAAACTTGTGAGGGGATGAACTCAAAAAGTTCAAGATGAAGCTGCTGACAGTGCAACTGCG
CTTTTGAACAGTCCCCTACTTGAGTTTTTCAAGTTCTACTTCGACGACTGTCACGTTGACGC

13 E N L S G D E L K K F K M K L L T V Q L R

187 AGAAGGCTATGGGCGCATCCACGCGGGGCCCCCTGCTGCAGATGGACGCCATAGATCTCACTG
TCTTCCGATACCCCGGTAGGGTGCGCCCGGGACGACGCTCTACCTGCGGTATCTAGAGTGAC

33 E G Y G R I P R G A L L Q M D A I D L T

249 ACAAACCTTGTGAGTACTATCTGGAGTCGTATGGCTTGGAGCTCACAATGACTGTGCTTAGA
TGTTTGAACAGTCGATGATAGACCTCAGCATACCGAACCTCGAGTGTTACTGACACGAATCT

54 D K L V S Y Y L E S Y G L E L T M T V L R

311 GACATGGGCTTACAGGAGCTGGCTGAGCAGCTGCAAACCGACTAAAGAAGAGTCTGGAGCTGT
CTGTACCCGAATGTCTCGACCGACTCGTCGACGTTTGCTGATTTCTTCTCAGACCTCGACA

75 D M G L Q E L A E Q L Q T T K E E S G A V

373 GGCAGCTGCAGCCAGTGTCCCTGCTCAGAGTACAGCCAGAACAGGACACTTTGTGGACCAGC
CCGTGACGTCGGTCACAGGGACGAGTCTCATGTTCGGTCTTGTCCTGTGAAACACCTGGTCG

95 A A A A S V P A Q S T A R T G H F V D Q

435 ACAGGCAAGCACTCATTGCCAGGGTCACAGAAGTGGACGGAGTGCTGGATGCTTTGTCATGGC
TGTCGGTTTCGTAGTAACGGTCCCAGTGTCTTCACCTGCCTCAGGACCTACGAAACGTACCG

116 H R Q A L I A R V T E V D G V L D A L H G

497 AGTGTGCTGACTGAAGGACAGTACCAGGCAGTTCGTGCAGAGACCACCAGCCAAGACAAGAT
TCACACGACTGACTTCCTGTTCATGGTCCGTCAAGCACGTCTCTGGTGGTGGTTCGTTCTA

137 S V L T E G Q Y Q A V R A E T T S Q D K M

559 GAGGAAGCTCTTCAGCTTTGTTCCATCCTGGAACCTGACCTGCAAGGACTCCCTCCTCCAGG
CTCCTTCGAGAAGTCGAAACAAGGTAGGACCTTGGAAGTGGACGTTCTGAGGGAGGAGGTCC

157 R K L F S F V P S W N L T C K D S L L Q

621 CCTTGAAGGAAATACATCCCTACTTGGTGATGGACCTGGAGCAGAGCTGAGGTATCTTTTCC
GGAACCTCCTTTATGTAGGGATGAACCACTACCTGGACCTCGTCTCGACTCCATAGAAAAGG (SEQ ID NO:3)

178 A L K E I H P Y L V M D L E Q S (SEQ ID NO:2)

683 AGCTACATTATCTAGCTCCTGACTTTGTATACACAATTTTTGAAAAACAATTTGTATTTGT
TCGATGTAATAGATCGAGGACTGAAACATATGTGTTAAAAACTTTTTTGTAAACATAAACA

745 GTTTAAAAAAGGGCGGCCGC (SEQ ID NO:1)
CAAAATTTTTTTTTTTTTTTTTTCCCGCCGGCG

FIG. 1

Applicant(s): John Bertin

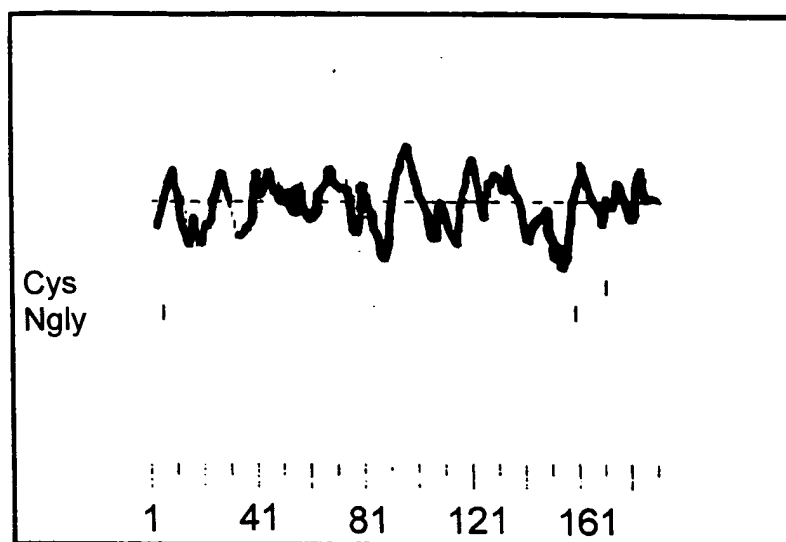
NOVEL MOLECULES OF THE CARD-RELATED PROTEIN
FAMILY AND USES THEREOF

FIG. 2

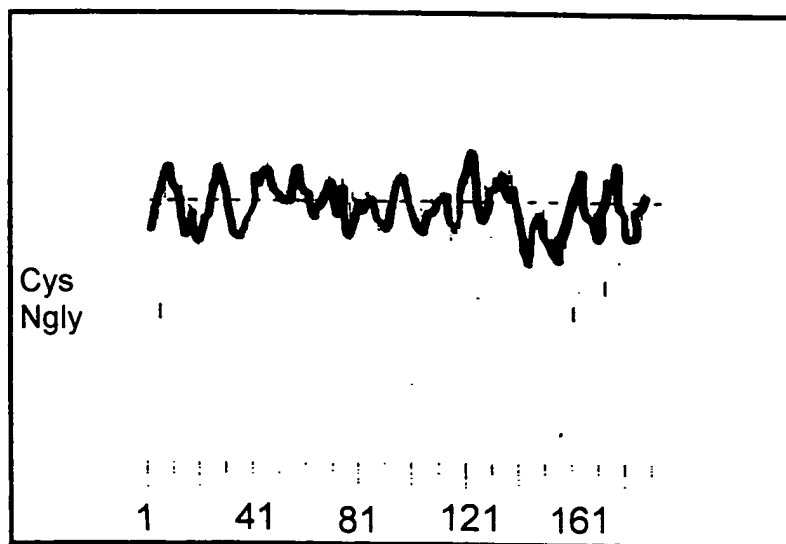


FIG. 4

Applicant(s): John Bertin

NOVEL MOLECULES OF THE CARD-RELATED PROTEIN
FAMILY AND USES THEREOF

1 CGCGTCCGGCTGCAGCGGGGTGAGCGGCGGCAGCGGCCGGGGATCCTGGAGCCATGGGGC
GCGCAGGCGCGACGTGCGCCCCACTGCGCGCCGTGCGCGGCCCTAGGACCTCGGTACCCCG
1► M G

61 GCGCGCGCGACGCCATCCTGGATGCGCTGGAGAACCTGACCGCCGAGGAGCTCAAGAAGT
CGCGCGCGCTGCGGTAGGACCTACGCGACCTCTTGGACTGGCGGCTCCTCGAGTTCTTCA
3► R A R D A I L D A L E N L T A E E L K K

121 TCAAGCTGAAGCTGCTGTGCGGTGCGGCTGCGCGAGGGCTACGGGCGCATCCCGCGGGGCG
AGTTGCACTTCGACGACAGCCACGGCGACGCGCTCCCGATGCCCCGCTAGGGCGCCCCCG
23► F K L K L L S V P L R E G Y G R I P R G

181 CGCTGCTGTCCATGGACGCCTTGGACCTCACCGACAAGCTGGTTCAGCTTCTACCTGGAGA
GCGACGACAGGTACCTGCGGAACCTGGAGTGGCTGTTGACCGAGTCAAGATGGACCTCT
43► A L L S M D A L D L T D K L V S F Y L E

241 CCTACGGCGCCGAGCTCACCGCTAACGTGCTGCGCGACATGGGCCTGCAGGAGATGGCCG
GGATGCCGCGGCTCGAGTGGCGATTGCACGACGCGCTGTACCCGACGTCTCTACCGGC
63► T Y G A E L T A N V L R D M G L Q E M A

301 GGCAGCTGCAGGCGGCCACGCACCAGGGCTCTGGAGCCGCGCCAGCTGGGATCCAGGCCC
CCGTCGACGTCCGCGGTTGCGTGGTCCCGAGACCTCGCGCGGTCGACCCCTAGGTCCGG
83► G Q L Q A A T H Q G S G A A P A G I Q A

361 CTCCTCAGTCGGCAGCCAAGCCAGGCCTGCACTTTATAGACCAGCACCGGGCTGCGCTTA
GAGGAGTCAGCCGTGCGTTTCGGTCCGGACGTGAAATATCTGGTCTGGCCCCGACGCGAAT
103► P P Q S A A K P G L H F I D Q H R A A L

421 TCGCGAGGGTCACAAACGTTGAGTGGCTGCTGGATGCTCTGTACGGGAAGGTCCTGACGG
AGCGCTCCAGTGTTTGCAACTCACCGACGACCTACGAGACATGCCCTTCCAGGACTGCC
123► I A R V T N V E W L L D A L Y G K V L T

481 ATGAGCAGTACCAGGCAGTGCAGGCGGAGCCACCAACCCAAGCAAGATGCGGAAGCTCT
TACTCGTCATGGTCCGTACGCCCCGGCTCGGGTGGTTGGGTTCTGTTCTACGCCTTCGAGA
143► D E Q Y Q A V R A E P T N P S K M R K L

541 TCAGTTTCACACCAGCCTGGAACCTGGACCTGCAAGGACTTGCTCCTCCAGGCCCTAAGGG
AGTCAAAGTGTGGTCCGACCTTGACCTGGACGTTCTGAACGAGGAGGTCCGGGATTCCC
163► F S F T P A W N W T C K D L L L Q A L R

601 AGTCCCAGTCCTACCTGGTGGAGGACCTGGAGCGGAGCTGAGGCTCCTTCCCAGCAACAC
TCAGGGTCAGGATGGACCACCTCCTGGACCTCGCCTCGACTCCGAGGAAGGGTCGTTGTG (SEQ ID NO:6)
183► E S Q S Y L V E D L E R S (SEQ ID NO:5)

661 TCCGGTCAGCCCCTGGCAATCCCAACCAATCATCCTGAATCTGATCTTTTTATACACAAT
AGGCCAGTCGGGGACCGTTAGGGTGGTTTAGTAGGACTTAGACTAGAAAAATATGTGTTA

721 ATACGAAAAGCCAGCTTGAA (SEQ ID NO:4)
TATGCTTTTCGGTCAACTT

FIG. 3

Applicant(s): John Bertin

NOVEL MOLECULES OF THE CARD-RELATED PROTEIN
FAMILY AND USES THEREOF

ALIGN calculates a global alignment of two sequences

version 2.0uPlease cite: Myers and Miller, CABIOS (1989)

> hCARD5-DNA

740 aa vs.

> mCARD5-DNA

763 aa

scoring matrix: paml20.mat, gap penalties: -12/-4

68.2% identity; Global alignment score: 2377

```

          10          20          30
inputs C---GCGTCCGGCTGCAG-CGGGGTG-----AGCG-GCGGCAGC-----GGC
      :  ::::::::::: :::: :  :::: ::::::::::: :::
      CCACGCGTCCGGCAGCAGGCAGGCTGCAGCAGGCGAGCAGCAGCAAGAGTAAAAGGTGAC
          10          20          30          40          50          60

          40          50          60          70          80          90
inputs CGGGGAT-----CCTGGAGCCATGGGGCGCGCGCGACGCCATCCTGGATGCGCTGGA
      :: :: :  :: ::::::::::: ::::: :  ::::::::::: :::: :
      CGCGGCTGCCCCACCCAGAGCCATGGGGCGGGCAGAGATGCCATCCTGGACGCTCTTGA
          70          80          90          100         110         120

          100         110         120         130         140         150
inputs GAACCTGACCGCCGAGGAGCTCAAGAAGTTCAAGCTGAAGCTGCTGTCGGTGCCGCTGCG
      :::: :::: :  :: ::::::::::: ::::::::::: ::::::::::: :::::
      AAACCTGTCAGGGGATGAACTCAAAAAGTTCAAGATGAAGCTGCTGACAGTGCAACTGCG
          130         140         150         160         170         180

          160         170         180         190         200         210
inputs CGAGGGCTACGGGCGCATCCCGCGGGGCGCGCTGCTGTCCATGGACGCCTTGGACCTCAC
      ::::::::::: ::::::::::: :  :: :::::  ::::::::::: :::::
      AGAAGGCTATGGGCGCATCCACGCGGGGCCCTGCTGCAGATGGACGCCATAGATCTCAC
          190         200         210         220         230         240

          220         230         240         250         260         270
inputs CGACAAGCTGGTCAGCTTCTACCTGGAGACCTACGGCGCCGAGCTCACCAGCTAAC-GTGC
      ::::::::::: ::::::::::: ::::::::::: :  ::  ::::::::::: . :::: ::::
      TGACAAACTTGTCTAGCTACTATCTGGAGTCGTATGGCTTGGAGCTCAC-AATGACTGTGC
          250         260         270         280         290

          280         290         300         310         320         330
inputs TGC GCGACATGGGCCTGCAGGAGATGGCCGGGCAGCTGCAGGCGGCCACGCACCAGGGCT
      :  ::::::::::: ::::::::::: :  ::::::::::: :  :  :::::
      TTAGAGACATGGGCTTACAGGAGCTGGCTGAGCAGCTGCAAACGACTAAAGA--AGAG-T
          300         310         320         330         340         350

          340         350         360         370         380         390
inputs CTGGAGCCGCGCCAGCTGGGATCCAGGCCCTCCTCAGTCGGCAGCCAAGCCAGGCTGTC
      ::::::::::: :  ::::: .. :  :  ::::: :::::  ::::::::::: :::
      CTGGAGCTGTGGCAGCTGCAGCCAGTGTCCCTGCTCAGAGTACAGCCAGAACAGG---AC
          360         370         380         390         400         410

          400         410         420         430         440         450

```

FIG. 5A

```
inputs ACTTTATAGACCACGACCGGGCTTATCGCGAGGGTCACAAACGTGAGTGGCCTGC  
::: :: : : : : : : : : : : : :  
ACTTTGTGGACCAGCACAGGCCAAGCACTCATTGCCAGGGTCACAGAAGTGACGGAGTGC  
420 430 440 450 460 470
```

460 470 480 490 500 510

```
inputs TGGATGCTCTGTACGGGAAGGTCCTGACGGATGAGCAGTACCAGGCAGTGCGGGCCGAGC  
::: :: : : : : : : : : : : : :  
TGGATGCTTTGCATGGCAGTGTGCTGACTGAAGGACAGTACCAGGCAGTTCGTGCAGAGA  
480 490 500 510 520 530
```

520 530 540 550 560 570

```
inputs CCACCAACCCAAGCAAGATGCGGAAGCTCTTCAGTTTCACACCAGCCTGGAACCTGGACCT  
::: :: : : : : : : : : : : : :  
CCACCAGCCAAGACAAGATGAGGAAGCTCTTCAGCTTTGTTCCATCCTGGAACCTGACCT  
540 550 560 570 580 590
```

580 590 600 610 620 630

```
inputs GCAAGGACTTGCTCCTCCAGGCCCTAAGGGAGTCCCAGTCCTACCTGGTGGAGGACCTGG  
::: :: : : : : : : : : : : : :  
GCAAGGACTCCCTCCTCCAGGCCCTTGAAGGAAATACATCCCTACTTGGTGATGGACCTGG  
600 610 620 630 640 650
```

640 650 660 670 680

```
inputs AGCGGAGCTGAGGC-TCCTTCCCAGCAACACTCCGGTC-AGCCCCCTGGCAAT-CCCAC-C  
::: :: : : : : : : : : : : : :  
AGCAGAGCTGAGGTATCTTTTTCCAGCTACATT---ATCTAGCTCCTGACTTTGTATAACAC  
660 670 680 690 700 710
```

690 700 710 720 730 740

```
inputs AAATCATCCTGAATCTGATCTTTTTATACACAATATACGAAAAACCAGCTTGAA (SEQ ID NO:4)  
::: :: : : : : : : : : : : : :  
AATTTTTGAAAAACAATT-TGTATTTGTGTTTAaaaaaaaaaaaaaaagg (SEQ ID NO:1)
```

720 730 740 750 760

FIG. 5B

Applicant(s): John Bertin

NOVEL MOLECULES OF THE CARD-RELATED PROTEIN
FAMILY AND USES THEREOF

ALIGN calculates a global alignment of two sequences

version 2.0uPlease cite: Myers and Miller, CABIOS (1989)

> hCARD5-protein 195 aa vs.

> mCARD5-protein 193 aa

scoring matrix: paml20.mat, gap penalties: -12/-4

71.8% identity; Global alignment score: 712

```

          10          20          30          40          50          60
inputs MGRARDAILDALENLTAEELKKFKLKLLSVPLREGYGRIPRGALLSMDALDLTDKLVSFY
       .....
       MGRARDAILDAENLSGDELKKFKMKLLTVQLREGYGRIPRGALLQMDAIDLTDKLVSY
          10          20          30          40          50          60

          70          80          90          100          110          120
inputs LETYGAELTANVLRDMGLQEMAGQLQAATHQGSGAAPAGIQAPPQSAAKPGLHFIQHRA
       .....
       LESYGLELTMTVLRDMGLQELAEQLQT-TKEESGAVAAAASVPAQSTARTG-HFVDQHRQ
          70          80          90          100          110

          130          140          150          160          170          180
inputs ALIARVTNVEWLLDALYGKVLTDQYQAVRAEPTNPSKMRKLFSFTPAWNWTCKDLLLQA
       .....
       ALIARVTEVDGVLDALHGSLVTEGQYQAVRAETTSQDKMRKLFSFVPSWNLTCKDSLLQA
       120          130          140          150          160          170

          190
inputs LRESQSYLVEDLERS (SEQ ID NO:5)
       .....
       LKEIHPYLVMDLEQS (SEQ ID NO:2)
       180          190

```

FIG. 6

Applicant(s): John Bertin

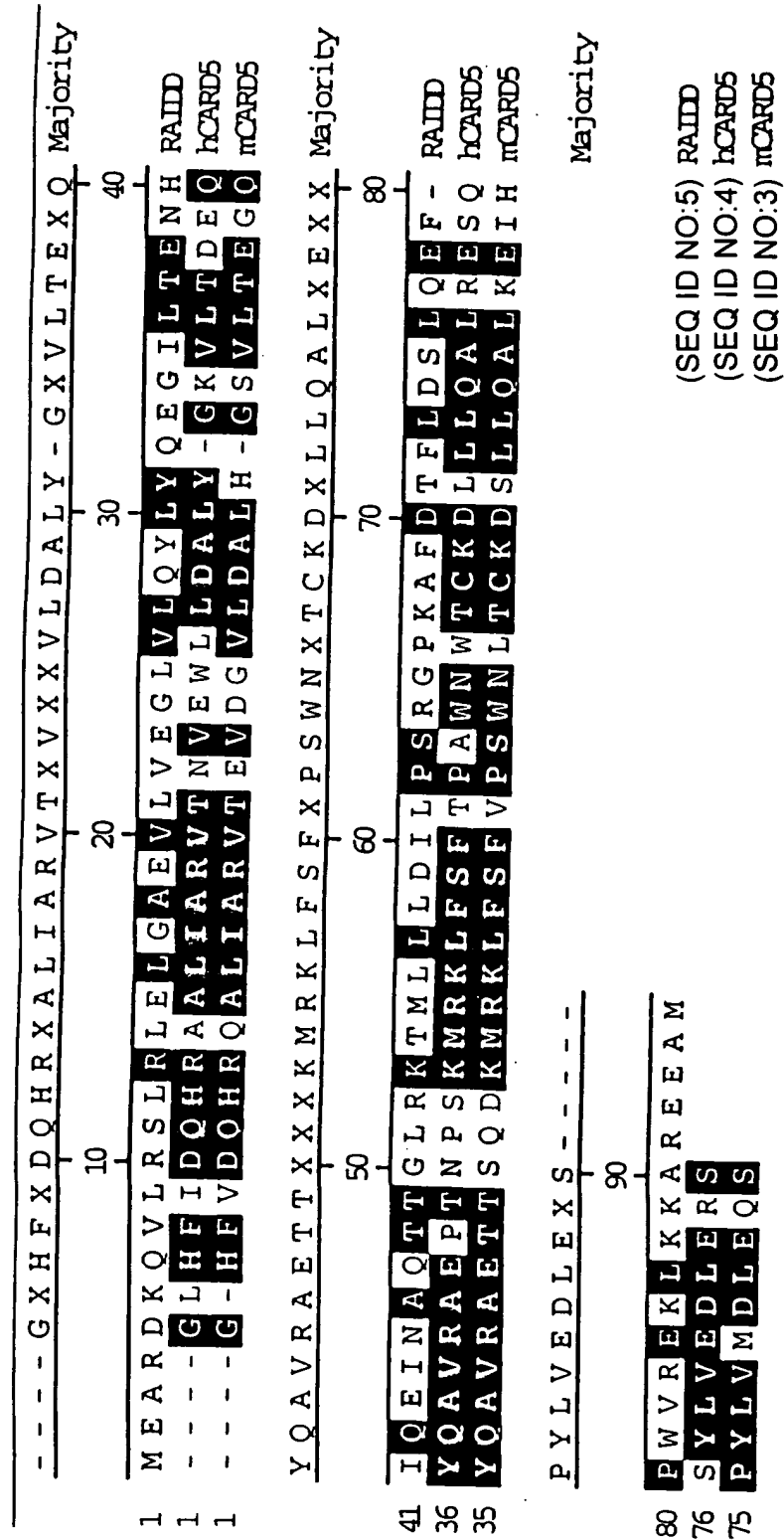
NOVEL MOLECULES OF THE CARD-RELATED PROTEIN
FAMILY AND USES THEREOF

FIG. 7

Applicant(s): John Bertin

NOVEL MOLECULES OF THE CARD-RELATED PROTEIN
FAMILY AND USES THEREOF

	-	A	-	E	S	-	G	S	E	I	I	D	Q	H	R	X	A	L	L	A	R	V	T	E	D	P	-	D	S	L	L	D	A	L	L	S	R	D	L	I	Majority	
		10		20		30		40																																		
1	I	A	Q	Q	W	-	-	-	-	I	Q	S	K	R	E	D	I	V	N	Q	M	T	E	A	C	L	N	Q	S	L	D	A	L	L	S	R	D	L	I	hCARD3-CARD		
1	-	-	-	-	E	S	H	P	H	I	Q	L	L	K	S	N	R	E	L	L	V	T	H	I	R	N	T	-	Q	C	L	V	D	N	L	L	K	N	D	Y	F	hCARD4-CARD
1	-	-	-	-	-	-	G	L	H	F	I	D	Q	H	R	A	A	L	I	A	R	V	T	N	-	-	V	E	W	L	L	D	A	L	L	Y	G	K	-	V	L	hCARD5-CARD
1	-	-	-	-	-	-	G	-	H	F	V	D	O	H	R	Q	A	L	I	A	R	V	T	E	-	-	V	D	G	V	L	D	A	L	L	H	G	S	-	V	L	hCARD5-CARD
1	M	A	T	E	S	T	P	S	E	I	I	E	R	E	R	K	K	L	L	E	I	L	Q	H	D	P	-	D	S	I	L	D	T	L	T	S	R	R	L	I	hCARD6-CARD	
1	M	A	S	E	G	A	S	S	E	I	I	E	K	Q	R	T	K	L	L	S	V	L	Q	Q	D	P	-	D	S	I	L	D	T	L	T	S	R	R	L	I	hCARD6-CARD	
	S	E	E	D	Y	E	A	V	E	A	E	T	T	X	L	S	K	V	R	K	L	L	I	L	V	Q	S	K	G	E	E	T	C	K	-	F	L	K	C	L	Majority	
		50		60		70		80																																		
36	M	K	E	D	Y	E	L	V	S	T	K	P	T	R	T	S	K	V	R	Q	L	L	D	T	T	D	I	Q	G	E	E	-	-	-	-	I	A	F	V	I	hCARD3-CARD	
37	S	A	E	D	A	F	I	V	C	A	C	P	T	Q	P	D	K	V	R	K	I	L	D	L	V	Q	S	K	G	E	E	V	S	E	F	F	I	-	Y	L	hCARD4-CARD	
32	T	D	E	Q	Y	Q	A	V	R	A	E	P	T	N	P	S	K	M	R	K	L	F	S	F	T	P	A	W	-	N	W	T	C	K	-	-	-	D	L	L	hCARD5-CARD	
31	T	E	G	Q	Y	Q	A	V	R	A	E	T	S	Q	D	F	M	R	K	L	F	S	F	V	P	S	W	-	N	L	T	C	K	-	-	-	D	S	L	hCARD5-CARD		
40	S	E	E	E	Y	E	T	L	E	N	V	T	D	L	L	K	K	S	R	K	L	L	I	L	V	Q	K	K	G	E	A	T	C	Q	H	F	L	K	C	L	hCARD6-CARD	
40	S	E	E	E	Y	E	T	L	E	A	I	T	D	P	L	K	K	S	R	K	L	L	I	L	I	C	K	K	G	E	D	S	C	C	F	L	K	C	L	hCARD6-CARD		
	L	Q	A	L	K	D	S	A	A	Y	L	G	L	D	P	E	V	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	Majority		
		90		100																																						
72	V	Q	K	L	K	D	N	K	Q	-	M	G	L	Q	P	Y	P	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	hCARD3-CARD		
76	L	Q	Q	L	A	L	-	-	A	Y	V	D	L	R	P	W	L	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	hCARD4-CARD		
68	L	Q	A	L	R	E	S	Q	S	Y	L	V	E	T	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	hCARD5-CARD			
67	L	Q	A	L	K	E	I	H	P	Y	L	V	M	E	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	hCARD5-CARD			
80	F	S	T	F	P	Q	L	A	I	C	G	L	R	H	E	V	L	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	-	hCARD6-CARD			
80	S	N	A	F	P	Q	S	A	S	T	L	G	L	K	Q	E	V	P	R	Q	G	T	G	E	V	F	V	S	-	-	-	-	-	-	-	-	-	-	hCARD6-CARD			

FIG. 8